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Strong approximations of level exceedences related to multiple hypothesis testing

PETER HALL¹ and QIYING WANG²

Particularly in genomics, but also in other fields, it has become commonplace to undertake highly multiple Student's t-tests based on relatively small sample sizes. The literature on this topic is continually expanding, but the main approaches used to control the family-wise error rate and false discovery rate are still based on the assumption that the tests are independent. The independence condition is known to be false at the level of the joint distributions of the test statistics, but that does not necessarily mean, for the small significance levels involved in highly multiple hypothesis testing, that the assumption leads to major errors. In this paper, we give conditions under which the assumption of independence is valid. Specifically, we derive a strong approximation that closely links the level exceedences of a dependent "studentized process" to those of a process of independent random variables. Via this connection, it can be seen that in high-dimensional, low sample-size cases, provided the sample size diverges faster than the logarithm of the number of tests, the assumption of independent t-tests is often justified.

Keywords: false discovery rate; family-wise error rate; genomic data; large deviation probability; moving average; Poisson approximation; Student's t-statistic; upper tail dependence; upper tail independence

1. Introduction

Today it is commonplace to undertake highly multiple hypothesis testing, generally in genomics and very often using tests based on Student's t-statistic; see, for example, Benjamini and Yekutieli (2001), Efron and Tibshirani (2002), Cui and Churchill (2003), Amaratunga and Cabrera (2004), page 114, Scheid and Spang (2005), Shaffer (2005), Fox and Dimmic (2006), Hu and Willsky (2006), Qiu and Yakovlev (2006), Efron (2007a), Liu and Hwang (2007) and van de Wiel and Kim (2007). This popularity of multiple t-testing also extends to other fields (e.g., Pawluk-Kołc et al. (2006)). The principal methods used to control the family-wise error rate and false discovery rate are founded on the assumption of independence among tests. Alternative approaches are generally

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¹Department of Mathematics and Statistics, The University of Melbourne, VIC 3010, Australia. E-mail: halpstat@ms.unimelb.edu.au

²School of Mathematics and Statistics F07, University of Sydney, NSW 2006, Australia. E-mail: qiyinq@maths.usyd.edu.au

based either on Bonferroni bounds, which are unsatisfactory for a variety of reasons (see, e.g., Perneger (1998)), or on the hope that, despite ample evidence of non-independence in terms of correlation analysis, independence can be assumed in practice.

The latter hope tends to be pinned either on work of Benjamini and Yekutieli (2001), who argued that in some settings, the absence of independence can give conservative results, or on experience with the analysis of financial data, which suggests that in some circumstances, it might be reasonable to assume that the upper tails of the test statistics are independent, even if the joint distributions are not. Upper tail independence, as it is sometimes called (for discussion, see, e.g., Wu (1994), Falk and Reiss (2001), R. Schmidt (2002), Li (2006), R. Schmidt and Stadtmüller (2006), T. Schmidt (2007)), is generally assumed to be non-asymptotic in nature. That is, tails of joint distributions are often taken to be perfectly independent beyond a certain threshold.

However, this type of model is not really appropriate for the analysis of genomic data. In particular, it is difficult to determine a biological reason for, or the actual location of, a threshold. It is of greater practical interest to consider the possibility that the strength of dependence in upper tails could become successively weaker as the number of simultaneous tests, and the number of data vectors, increases. If this could be established in the context of tests based on Student's t-statistic, it would lend immediate justification to the often-made assumption (see the articles cited in the first paragraph of this paper) that highly multiple t-statistics can be taken to be independent.

The present paper will establish such a result. The mechanism for our model involves the critical points for tests becoming more extreme as the number, p, of tests diverges (in fact, the increase in critical points is a direct consequence of p diverging) so that the tests are conducted further into the tails; furthermore, the tails of the distributions of test statistics becoming successively lighter as the number of degrees of freedom of the test statistics increases.

We impose particularly weak conditions on the marginal distributions of components. In particular, the distributions need only three finite moments. With this assumption, and permitting the size, n, of the group sample to increase a little faster than the logarithm of the number of tests, it follows from our results that the joint distributions of test statistics enjoy an asymptotic form of the upper tail independence property.

This result would not be so striking if the statistics had normal distributions, but it fails for heavy-tailed distributions such as those for which not all moments are finite. Of course, Student's t-distribution is itself in this category, yet our results show that asymptotic independence holds in a particularly strong sense for Student's t-statistic, even if it is computed from relatively heavy-tailed data. The reason this is possible is that we permit the group sample size to increase at a rate that is just sufficient to convert heavy tails to tails that are sufficiently light, to enable approximate independence at high levels.

It can be seen from this property that the availability of upper-tail asymptotic independence is a bonus of working with highly multiple hypothesis testing, that is, with "large p and small n" problems. It is not available in more conventional, "small p and large n" problems, where there is a very large literature on modelling dependence in highly multiple hypothesis testing.

There is a literature on comparing studentized means when the variances used for studentizing are computed from pooled data and so are common to each test statistic. However, in our experience, that approach is used less frequently, in practice, than the "local" standardization treated in the present paper. When using the latter method, each mean is divided by the standard deviation of the sample from which it was computed. A major motivation is that the true variances may be different in each instance. Even if the variances can reasonably be assumed to be the same, it can be desirable to use the local approach since it confers greater robustness. For example, when applied to the mean alone, rather than its locally studentized form, the large-deviation properties that underpin the analysis of high-level exceedences require the data to have lighter tails.

Statistical literature on highly multiple hypothesis testing is outlined in helpful reviews by Hochberg and Tamhane (1987), Pigeot (2000), Dudoit et al. (2003), Bernhard et al. (2004) and Lehmann and Romano (2005), Chapter 9. Benjamini and Hochberg (1995) introduced an approach, which has become very popular, to the controlling of false discovery rates; see also Simes (1986), Hommel (1988), Hochberg (1988), Sarkar and Chang (1997), Sarkar (1998), Sen (1999), Hochberg and Benjamini (1990) and Lehmann et al. (2005). Benjamini and Yekutieli (2001) specified conditions under which simultaneous, dependent hypothesis tests, conducted as though they were independent, give conservative results; Benjamini and Yekutieli (2005) addressed similar issues in the context of false coverage-statement rate. Sarkar (2002) extended the work of Benjamini and Yekutieli (2001). Efron (2007b) suggested correlation corrections for large-scale simultaneous hypothesis testing. Blair et al. (1996) proposed methods for controlling family-wise error rates in multiple procedures, Holland and Cheung (2002) discussed robustness of family-wise error rates and Clarke and Hall (2009) discussed robustness of testing procedures based on means.

2. Results and applications

2.1. Model and main results

Given $p, n \geq 1$, assume that for $1 \leq i \leq p$ and $1 \leq j \leq n$, we observe data U_{ij} , which we use to construct t-statistics $T_i = n^{1/2} \bar{U}_i / S_i$, where $\bar{U}_i = n^{-1} \sum_j U_{ij}$ and $S_i^2 = n^{-1} \sum_j U_{ij}^2 - \bar{U}_i^2$. In practice, the statistic T_i is used to test the hypothesis that the ith group has zero mean, against a one-sided alternative. When controlling the level of family-wise error rate (FWER) for step-down tests, we require the values of probabilities $P(T_i > t \text{ for } i = i_1, \ldots, i_k)$ for different levels t and different subsets $\{i_1, \ldots, i_k\}$ of $\{1, \ldots, p\}$. Theorem 1 below will enable us to compute these through approximation by the case where the T_i 's are all independent; see Section 2.3 for further details.

We standardize S_i^2 by dividing by n, rather than n-1, since the former is more common in nonparametric problems, but the results below are unaffected by this issue. Since we studentize, there is no loss of generality in assuming that the variance of each

component equals 1. More particularly, we ask that

$$0 \le E(U_{i1}) = d_i, \quad \text{var}(U_{i1}) = 1 \quad \text{for all } i, \quad \sup_{i \ge 1} E(|U_{i1}|^3) < \infty, \quad (2.1)$$

where d_1, d_2, \ldots is a sequence of constants. The assumption that $d_i \geq 0$ is made here because, in the great majority of practical applications, the hypothesis alternative to the null entails the zero level being exceeded. Accordingly, the tests are one-sided, hence our preoccupation with exceedences of a level. However, minor modifications of our arguments permit the two-sided case to be treated.

Further, we assume that for an integer $\kappa > 0$,

the random vectors (U_{1i}, U_{2i}, \ldots) , for $i \geq 1$, are independent and identically distributed, the sequence of random variables U_{11}, U_{21}, \ldots is κ -dependent and $\max_{i_1, i_2: i_1 \neq i_2} \rho_{i_1 i_2} < 1$,

where $\rho_{i_1i_2} = \text{corr}(U_{i_11}, U_{i_21})$. The third moment condition in (2.1) permits the variables U_{ij} to have relatively heavy-tailed distributions, for example, a Pareto distribution with tail exponent greater than 3.

The assumption of short-range correlation in (2.2) is, of course, an oversimplification, but it reflects the low level of correlation that is often observed in practice. For example, Messer and Arndt (2006) argue that correlation decays from about 0.08, at a separation of approximately two base pairs, to about 0.01 for a separation of ten base pairs. Results reported by Mansilla et al. (2004) corroborate these figures if we assume that their data are normally distributed. More generally, Almirantis and Provata (1999) give evidence of both short-range and long-range correlation, depending on the nature of the DNA or RNA under investigation.

The relationship between the group size, n, and the number of hypothesis tests, p, is assumed to satisfy

$$\log p = o(n). \tag{2.3}$$

This allows the group size to be very much smaller than the number of tests. In the absence of more detailed assumptions about the distributions of the U_{ij} 's, (2.3) is necessary for the theorem we shall give below. To appreciate why, note that if the U_{ij} 's are independent and identically distributed with an atom at zero and, in particular, if $\delta \equiv P(U_{ij} = 0) > 0$, then, with probability at least δ^n , the t-statistic T_1 assumes the indeterminate value 0/0. In such cases, we shall take $T_1 = 1$, but in order for the theorem to have a meaningful interpretation when t is the $(1-p^{-1})$ -level quantile of the standard normal distribution, it is essential that the probability that $T_1 = 0/0$ be of smaller order than p^{-1} . Therefore, we require $\delta^n = o(p^{-1})$ for all $0 < \delta < 1$ and this assumption is equivalent to (2.3).

Define

$$\alpha = \frac{1}{4} \min_{i_1, i_2 : i_1 \neq i_2} (1 - \rho_{i_1 i_2})$$
(2.4)

and $\gamma = \alpha + 1$. Condition (2.2) implies that $\alpha > 0$ and, of course, $\alpha \le \frac{1}{2}$. Given $\eta > 0$, let t = t(p) satisfy

$$(1+\eta)\sqrt{2\gamma^{-1}\log p} \le t = \mathcal{O}(\sqrt{\log p}), \qquad \max_{1 \le i \le p} d_i = \mathcal{O}(t/\sqrt{n}), \tag{2.5}$$

where d_1, d_2, \ldots are as in (2.1). If t satisfies the first part of (2.5), then any function ϕ which satisfies, as $t \to \infty$,

$$\phi(t) = \exp\{o(t^2)\}\{\exp(-\frac{1}{4}t^2) + p\exp(-\gamma t^2/2)\}$$
(2.6)

converges to zero as $p \to \infty$. In the arguments in Section 3, we shall use this notation generically; while ϕ will satisfy (2.6), it will alter from one appearance to another. Strictly speaking, it is not essential to take p to diverge. Although that condition motivates the assumption of divergent t and is, in turn, motivated by the contemporary high-dimensional problems that led to this work, it is not necessary for the theorem below.

Theorem 1. If (2.2)–(2.6) hold, then there exists a probability space on which are defined random variables $T_1^{\text{new}}, \ldots, T_p^{\text{new}}$ and T_1', \ldots, T_p' such that (i) the joint distribution of $T_1^{\text{new}}, \ldots, T_p^{\text{new}}$ is identical to that of T_1, \ldots, T_p ; (ii) the random variables T_1', \ldots, T_p' are independent and distributed, respectively, as T_1, \ldots, T_p ; and (iii) with probability equal to $1 - \phi(t)$, the exceedences of t by $T_1^{\text{new}}, \ldots, T_p^{\text{new}}$ occur at the same indices and take the same values as the exceedences of t by T_1', \ldots, T_p' .

To interpret the theorem, note that we would normally expect the dependent data set T_1, \ldots, T_p to exhibit clusters of level exceedences, rather than the single, isolated exceedences associated with the independent sequence T'_1, \ldots, T'_p . The fact that the T_i process (or, equivalently, the T_i^{new} process) behaves like the T'_i process in the case of large exceedences reflects the fact that, since the marginal distribution of a t-statistic is relatively light-tailed (if n is sufficiently large – see (2.3)), exceedences of a high level are rare and so are unlikely to occur together. The case of low-level exceedences is a very different matter, of course, and so we would expect the theorem to fail if the lower bound for t, in the first part of (2.5), were relaxed too far.

2.2. Applications

In this section, we treat the case of the null hypothesis, where $d_i = 0$ for each i. This would be assumed in most applications of Theorem 1 since it represents the setting that is conventionally used for calibration.

The theorem implies that, in a strong sense, exceedences down to those of the level $(1+\eta)(2\gamma^{-1}\log p)^{1/2}$ are identical to the ones that would occur in the case of independent tests. Now, the probability associated with an exceedence of $(1+\eta)(2\gamma^{-1}\log p)^{1/2}$ is, for small η , approximately $p^{-1/\gamma}$. Therefore, false discoveries at probability levels of approximately $p^{-1/\gamma}$, and at lower levels, can be adequately controlled by assuming that

the tests are independent, even when they are not. Note that $\gamma^{-1} < 1$ and that the false-discovery level controlled by the conventional family-wise error rate is only p^{-1} .

Next, we discuss the sorts of calculations that are enabled by Theorem 1. Let Q_j denote the number of indices $i \in [1,p]$ for which T_i lies in the interval $(t_j,t_{j-1}]$, where $j \geq 1$ and t_j is determined by $P(T_1 > t_j) = j\beta/p$, with $\beta > 0$ held fixed. (We take $t_0 = \infty$.) If T_1, \ldots, T_p were fixed, then the joint distribution of the k+1 random variables $Q_1, \ldots, Q_k, p - \sum_j Q_j$ would be exactly multinomial with parameters p and $q_1, \ldots, q_k, 1 - \sum_j q_j$, where $q_j = P(T_1 \in (t_j, t_{j-1}])$. Theorem 1 implies that for the dependent process T_1, \ldots, T_p , and for any $k_0 = k_0(p)$ for which t_{k_0} satisfies (2.5), any simultaneous probability calculation based on the multinomial result, but applied to the actual T_j process rather than an idealized process with independent marginals, is valid, provided that $k \leq k_0$ and the final computed probability is quantified by adding an error which is stated to be of order $\exp\{o(t_k^2)\}\{\exp(-t_k^2/4) + p\exp(-\gamma t_k^2/2)\}$. The latter probability converges to zero, even if $k = k_0$ is taken as large as $p^{(\gamma_1 - 1)/\gamma_1}$, where $\gamma_1 \in (1, \gamma)$.

From this point, simultaneous multinomial probability calculations based on Q_1, \ldots, Q_k , familiar from the well-understood case of independent test statistics, can be used to construct rules for controlling FWER or false discovery rate (FDR); see, for example, Benjamini and Hochberg (1995). Wang and Hall (2009) have shown that, under the assumption of finite third moments, highly accurate approximations are available for the marginal distribution of T_1 . Such calculations, which justify standard normal, Student's t- or bootstrap approximations to the marginal distribution of T_1 , are already widely used in practice (see Section 1), in conjunction with the independence assumption, when controlling false discovery rates. Our paper provides justification for these methods.

More generally, Theorem 1 implies that if a probability statement about what the process T_1, \ldots, T_p does above the level t is founded on the assumption of independence, then, no matter how complex or convoluted the statement might be, the claimed probability level is accurate to within $\phi(t)$.

To give an example of calculations based on Theorem 1, take $p \le p_0 = 10^6$, n = 100 and t = 5.052, the latter denoting the upper $(1 - p_0^{-1})$ -level quantile of Student's t-distribution with n - 1 = 99 degrees of freedom. Reflecting empirical evidence given in Section 2.1, take $\gamma = \frac{1}{4}(1-0.1)+1=1.225$. Then, (2.5) is in order; the probability that at least one value of p independent t statistics, each on 99 degrees of freedom, exceeds t = 5.052 equals 0.010, 0.095 and 0.63 for $p = 10^4$, 10^5 and 10^6 , respectively; and (2.6) suggests that the errors in these levels are in error by less than 30%, 20% and 0.25%, respectively. Most likely, the errors are much less than these since the asymptotic bound is derived only as an upper bound. If we were to make a general probability statement about exceedences of the level 5.052 by the stochastic process of t statistics, under the assumption of independence, then, despite the process actually being κ -dependent rather than independent, we would expect to make errors no greater than these respective values. In the same general setting, relative error decreases to zero as p and t increase. For example, in cases where t solves $1 - \Phi(t) \approx p^{-1}$, with Φ denoting the standard normal distribution function, we have $\exp(-\frac{1}{4}t^2) + p \exp(-\frac{1}{2}\gamma t^2)\}/(1-p)^p = O[\exp(-\frac{1}{4}t^2) + t \exp\{-\frac{1}{2}(\gamma - 1)t^2\}] \to 0$ as $t \to \infty$.

2.3. Generalizations

Theorem 1 can be extended to other settings, in particular, to those where (a) a wider range of dependence, obtained by allowing κ in (2.2) to diverge with p, is allowed; (b) the value of n for the ith group equals n_i , depending on i, and (2.3) is altered by requiring that $\log p = \mathrm{o}(\min_{i \leq p} n_i)$; (c) weights w_{ij} are incorporated into the construction of the t-statistics T_i , by defining $\bar{U}_i = n_i^{-1} \sum_j w_{ij} U_{ij}$, $S_i^2 = n_i^{-1} \sum_j w_{ij}^2 U_{ij}^2 - \bar{U}_i^2$ and, as before, $T_i = n_i^{1/2} \bar{U}_i / S_i$. Provided the weights satisfy

$$\sup_{i,j} |w_{ij}| \le C_1, \qquad \inf_{1 \le i \le p} n_i^{-1} \# \{j \colon |w_{ij}| \ge C_2\} \ge C_3,$$

where C_1, C_2, C_3 are positive constants not depending on p, the proof in this more general case is as in Section 3. However, the statement of the theorem is then less elegant and less transparent, so we do not give the more general version here. Incorporation of the weights w_{ij} permits the scope of the example above to be extended to hypothesis-testing problems involving linear regression.

To indicate the types of results that can be achieved under longer ranges of dependence, we shall discuss the case of a moving average,

$$U_{ij} = \kappa^{-1/2} \sum_{k=1}^{\kappa} \varepsilon_{j,i+k},$$

where $\kappa = \kappa(p)$ is permitted to diverge to infinity at a rate not exceeding $\log p$ and the independent disturbances ε_{ji} are all distributed as ε , for which $E(\varepsilon) = 0$ and $E|\varepsilon|^3 < \infty$. In this setting, (2.2) holds. We strengthen (2.3) by asking that $\log p = O(n^{1/3})$. The definition of t implicit in (2.5) can now be refined to

$$t = \sqrt{2\gamma^{-1}(\log p + A\log\log p)},$$

where A > 0 denotes a sufficiently large absolute constant. The conclusions of Theorem 1 continue to hold, with a similar proof if we replace $1 - \phi(t)$ by 1 - o(1).

3. Proof of Theorem 1

3.1. Step 1: Preliminaries

The notation $D_1, D_2, ...$ will denote constants not depending on n or p. Let $Q_i = n^{-1} \sum_j (U_{ij} - d_i)^2$, $R_i = n^{1/2} \bar{U}_i / Q_i^{1/2}$ and note that

for each
$$t > 0$$
, the events $T_i > t$ and $R_i > t/(1 + n^{-1}t^2)^{1/2}$ are identical. (3.1)

Also, note that
$$R_i = (\sum_i V_{ij} + nd_i)/(\sum_i V_{ij}^2)^{1/2}$$
, where $V_{ij} = U_{ij} - d_i$, $1 \le j \le n$, are

independent and identically distributed random variables satisfying

$$0 \le E(V_{i1}) = 0,$$
 $E(V_{i1}^2) = 1$ for all i , $\sup_{i>1} E(|V_{i1}|^3) < \infty$; (3.2a)

cf. (2.1).

3.2. Step 2: Probabilities of exceedences in ones and twos

Using results of Wang and Hall (2009) (see also Wang (2005)), it can be shown that, for constants $D_1, D_2, D_3 > 0$, and whenever $0 < s < D_1 n^{1/2}$,

$$P(R_i > s) \le D_2 s^{-1} \exp(D_3 s^3 n^{-1/2} - \frac{1}{2} s^2 + \sqrt{n} d_i s).$$
 (3.2)

We also wish to prove the following related result for pairs of exceedences.

Lemma. Assume the conditions of Theorem 1. There then exist $D_4, D_5 > 0$ such that for all i_1, i_2 with $i_1 \neq i_2$, and for all $0 < s < D_4 n^{1/2}$, we have

$$\sup_{1 \le |i_1 - i_2| \le k_2 - k_1} P(R_{i_1} > s, R_{i_2} > s)$$

$$\le 5 \exp\{-\frac{1}{2}(1 + \alpha)s^2 + D_5 n^{-1/2} s^3 + 2\sqrt{n}(d_{i_1} + d_{i_2})s\},$$
(3.3)

where α is as in (2.4).

To establish the lemma, we write

$$\begin{split} U_{i_{1}1}^{(1)} &= V_{i_{1}1}I(|V_{i_{1}1}| \leq n^{1/2}/s, |V_{i_{2}1}| \leq n^{1/2}/s), \qquad U_{i_{1}1}^{(2)} &= V_{i_{1}1} - U_{i_{1}1}^{(1)}, \\ U_{i_{2}1}^{(1)} &= V_{i_{2}1}I(|V_{i_{1}1}| \leq n^{1/2}/s, |V_{i_{2}1}| \leq n^{1/2}/s), \qquad U_{i_{2}1}^{(2)} &= V_{i_{2}1} - U_{i_{1}1}^{(1)}. \end{split}$$

By virtue of (3.2a), simple calculations show that

$$|E(U_{i_{1}1}^{(1)} + U_{i_{2}1}^{(1)})| \le D_{6}s^{2}/n,$$

$$E\{(U_{i_{1}1}^{(1)})^{2} + (U_{i_{2}1}^{(1)})^{2}\} = 2 + O(1)s/\sqrt{n},$$

$$E(U_{i_{1}1}^{(1)} + U_{i_{2}1}^{(1)})^{2} = E\{(V_{i_{1}1} + V_{i_{2}1}) - (U_{i_{1}1}^{(2)} + U_{i_{2}1}^{(2)})\}^{2}$$

$$\le 2(1 + \rho_{ij}) + D_{6}s/\sqrt{n},$$

$$E|U_{i_{1}1}^{(1)} + U_{i_{2}1}^{(1)}|^{3} \le D_{6}.$$

These results, and the bound $e^x \le 1 + x + \frac{1}{2}x^2 + \frac{1}{6}|x|^3e^x$ (valid for all real x), imply that, with $h = s/\sqrt{n}$,

$$E\left[\exp\left\{\frac{1}{2}h(V_{i_{1}1} + V_{i_{2}1}) - \frac{1}{4}h^{2}(V_{i_{1}1}^{2} + V_{i_{2}1}^{2})\right\}I(|V_{i_{1}1}| \le n^{1/2}/s, |V_{i_{2}1}| \le n^{1/2}/s)\right]$$

$$\le 1 + (\rho_{ij} - 1)\frac{s^{2}}{4n} + D_{7}\frac{s^{3}}{n^{3/2}},$$

$$E\left[\exp\left\{\frac{1}{2}h(V_{i_{1}1} + V_{i_{2}1}) - \frac{1}{4}h^{2}(V_{i_{1}1}^{2} + V_{i_{2}1}^{2})\right\}I(|V_{i_{1}1}| \ge n^{1/2}/s, \text{ or } |V_{i_{2}1}| \ge n^{1/2}/s)\right]$$

$$\le \exp\{P(|V_{i_{1}1}| > n^{1/2}/s) + P(|V_{i_{2}1}| > n^{1/2}/s)\}$$

$$< D_{8}(s/\sqrt{n})^{3}.$$

$$(3.5)$$

Results (3.4) and (3.5), together with the independence of V_{i_1k} for each i_1 , imply that, for $s \leq D_9 \sqrt{n}$, with D_9 sufficiently small,

$$E\left[\exp\left\{\frac{h}{2}\sum_{k}(V_{i_{1}k}+V_{i_{2}k})-\frac{h^{2}}{4}\sum_{k}(V_{i_{1}k}^{2}+V_{i_{2}k}^{2})\right\}\right]$$

$$\leq\left\{1+(\rho_{ij}-1)\frac{s^{2}}{4n}+D_{10}\frac{s^{2}}{n^{3/2}}\right\}^{n}\leq\exp\left\{(\rho_{ij}-1)\frac{s^{2}}{4}+D_{11}\frac{s^{3}}{\sqrt{n}}\right\}.$$
(3.6)

Define $\varepsilon^2 = (1 - \rho_{ij})/8$. It follows from (3.6) that whenever $s \leq D_9 \sqrt{n}$, with D_9 sufficiently small, we have

$$\pi_{1n} \equiv P \left\{ 2h \sum_{k} (V_{i_1k} + V_{i_2k}) - h^2 \sum_{k} (V_{i_1k}^2 + V_{i_2k}^2) + 2\sqrt{n} (d_{i_1} + d_{i_2}) s \ge 2s^2 (1 - \varepsilon^2) \right\}$$

$$\leq \exp \left\{ 2\sqrt{n} (d_{i_1} + d_{i_2}) s - \frac{1}{2} s^2 (1 - \varepsilon^2) \right\}$$

$$\times E \left[\exp \left\{ \frac{1}{2} h \sum_{k} (V_{i_1k} + V_{i_2k}) - \frac{1}{4} h^4 \sum_{k} (V_{i_1k}^2 + V_{i_2k}^2) \right\} \right]$$

$$\leq \exp \left\{ -\frac{1}{2} (1 + \alpha) s^2 + D_{12} \frac{s^3}{\sqrt{n}} + 2\sqrt{n} (d_{i_1} + d_{i_2}) s \right\},$$
(3.7)

where α is as defined in (2.4). Write $\Omega_n = (1 - \varepsilon, 1 + \varepsilon)$ and note that if $0 < \varepsilon < \frac{1}{2}$, then

$$\left\{ \sum_{k} V_{i_2k} + nd_{i_2} \ge s(nQ_{i_2}^2)^{1/2}, Q_{i_2} \in \Omega_n \right\}$$

$$\subseteq \left\{ 2h \sum_{k} V_{i_2k} - h^2 \sum_{k} V_{i_2k}^2 + 2\sqrt{n} d_{i_2} s \ge s^2 (1 - \varepsilon^2) \right\},$$

where $h = s/\sqrt{n}$. It can be shown that

$$P(R_{i_1} > s, R_{i_2} > s)$$

$$= P\left\{ \sum_{k} V_{i_1k} + nd_{i_1} \ge s(nQ_{i_1}^2)^{1/2}, \sum_{k} V_{i_2k} + nd_{i_2} \ge s(nQ_{i_2}^2)^{1/2} \right\}$$

$$\le \pi_{1n} + \pi_{2n} + \pi_{3n} + \pi_{4n} + \pi_{5n},$$
(3.8)

where

$$\begin{split} \pi_{2n} &= P \bigg\{ \sum_{k} V_{i_1k} + n d_{i_1} \geq s (nQ_{i_1}^2)^{1/2}, Q_{i_1} \geq 1 + \varepsilon \bigg\}, \\ \pi_{3n} &= P \bigg\{ \sum_{k} V_{i_2k} + n d_{i_2} \geq s (nQ_{i_2}^2)^{1/2}, Q_{i_2} \geq 1 + \varepsilon \bigg\}, \\ \pi_{4n} &= P \bigg\{ \sum_{k} V_{i_1k} + n d_{i_1} \geq s (nQ_{i_1}^2)^{1/2}, Q_{i_1} \leq 1 - \varepsilon \bigg\}, \\ \pi_{5n} &= P \bigg\{ \sum_{k} V_{i_2k} + n d_{i_2} \geq s (nQ_{i_2}^2)^{1/2}, Q_{i_2} \leq 1 - \varepsilon \bigg\}. \end{split}$$

Property (3.3) will follow from (3.7) and (3.8) if we prove that there exists $D_{13} > 0$ such that, for $s \le D_{13} n^{1/2}$,

$$\pi_{kn} \le \exp\left\{-\frac{1}{2}(1+\alpha)s^2 + D_{12}\frac{s^3}{\sqrt{n}} + 2\sqrt{n}(d_{i_1} + d_{i_2})s\right\}$$
(3.9)

for k = 2, 3, 4, 5.

Our proof of (3.9) is based on arguments of Shao (1999) (see also the proof of Proposition 4.2 of Wang and Hall (2009)) and uses the following result: if EX = 0, $EX^2 = 1$ and $E|X|^3 < \infty$, then for any $\lambda > 0$, $\theta > 0$ and x > 0,

$$E[\exp\{\lambda bX - \theta(bX)^2\}] = 1 + (\lambda^2 - \theta)n^{-1}x^2 + A(\lambda, \theta)n^{-3/2}x^3E|X|^3,$$
 (3.10)

where $b = x/\sqrt{n}$ and $A(\lambda, \theta)$ depends only on λ and θ . This result is a special case of Lemma 1 of Shao (1999). Also, note that

$$\pi_{2n} \le \pi_{2n}^{(1)} + P\left\{ \sum_{k} V_{i_1k} + nd_{i_1} \ge s(nQ_{i_1}^2)^{1/2}, Q_{i_1} \ge 3 \right\}$$

$$\le \pi_{2n}^{(1)} + \pi_{2n}^{(2)} + \pi_{2n}^{(3)},$$

where, noting that $\sqrt{n}|d_i| \leq s/5$, we define

$$\pi_{2n}^{(1)} = P\left\{\sum_{k} V_{i_1k} + nd_{i_1} \ge s(nQ_{i_1}^2)^{1/2}, 1 + \varepsilon \le Q_{i_1} < 3\right\},$$

$$\pi_{2n}^{(2)} = P\left\{\sum_{k} V_{i_1k} I(|V_{i_1k}| > n^{1/2}/s) \ge s\left(\sum_{k} V_{i_1k}^2\right)^{1/2}\right\},$$

$$\pi_{2n}^{(3)} = P\left\{\sum_{k} V_{i_1k} I(|V_{i_1k}| \le n^{1/2}/s) \ge 3s\sqrt{n}/2\right\}.$$

If the random variable H has the Bi(n,p) distribution and if a>0, then $P(H>an)\leq (ep/a)^{an}$ and so

$$\pi_{2n}^{(2)} \le P \left\{ \sum_{k} I(|V_{i_1k}| > n^{1/2}/s) \ge s^2 \right\}$$

$$\le \left\{ s^{-2} 12nP(|V_{i_1k}| > n^{1/2}/s) \right\}^{s^2} \le \frac{1}{2} e^{-s^2}$$

for $s \leq D_{14}\sqrt{n}$, with D_{14} sufficiently small. Arguments similar to those in the proof of (3.7) yield that $\pi_{2n}^{(3)} \leq \frac{1}{2} \mathrm{e}^{-s^2}$ for $s \leq D_{14}\sqrt{n}$ with D_{14} sufficiently small. To estimate $\pi_{2n}^{(1)}$, we write $\mathcal{S}_1 = \{(x,y)\colon x \geq s\sqrt{y}, s^2(1+\varepsilon)^2 \leq y \leq 9s^9\}$. It follows from (3.10) with $\lambda = 1$, $\theta = \frac{1}{6}$ and $X = V_{i_11}$ that, with $h = s/\sqrt{n}$,

$$\pi_{2n}^{(1)} = P\left\{ \left(h \sum_{k} V_{i_1k} + \sqrt{n} d_i s, h^2 \sum_{k} V_{i_1k}^2 \right) \in \mathcal{S}_1 \right\}$$

$$\leq E\left[\exp\left(h \sum_{k} V_{i_1k} - \frac{1}{6} h^2 \sum_{k} V_{i_1k}^2 + \sqrt{n} d_i s \right) \exp\left\{ - \inf_{(x,y) \in \mathcal{S}_1} (x - y/6) \right\} \right]$$

$$\leq \exp\left\{ \left(\frac{1}{2} - \frac{1}{6} \right) s^2 - s^2 (1 + \varepsilon) + \frac{1}{6} s^2 (1 + \varepsilon)^2 + \sqrt{n} d_i s + D_{15} s^3 n^{-1/2} \right\}$$

$$\leq \exp\left\{ - \frac{1}{2} s^2 - (5\varepsilon s^2/8) + \sqrt{n} d_i s + (D_{15} s^3/\sqrt{n}) \right\}$$

$$\leq \exp\left\{ - \frac{1}{2} (1 + \alpha) s^2 + \sqrt{n} d_i s + (D_{15} s^3/\sqrt{n}) \right\},$$

where we have used the fact that the function $f(y) = s\sqrt{y} - \frac{1}{6}y$ is increasing in $s^2(1+\varepsilon)^2 \le y \le 9s^2$. Combining all of the above estimates, we obtain

$$\pi_{2n} \le \exp\{-\frac{1}{2}(1+\alpha)s^2 + \sqrt{n}d_is + (D_{15}s^3/\sqrt{n})\}.$$

Similarly, we may prove (3.9) for k = 3.

Put $S_2 = \{(x,y): x \ge s\sqrt{y}, y \le (1-\varepsilon)^2 s^2\}$. It follows from (3.10) with $\lambda = 1$, $\theta = 2$ and $X = V_{i_1 1}$ that, with $h = s/\sqrt{n}$,

$$\pi_{4n} = P\left\{ \left(h \sum_{k} V_{i_1 k} + \sqrt{n} d_i s, h^2 \sum_{k} V_{i_1 k}^2 \right) \in \mathcal{S}_2 \right\}$$

$$\leq E\left[\exp\left(h \sum_{k} V_{i_1 k} - 2h^2 \sum_{k} V_{i_1 k}^2 + \sqrt{n} d_i s \right) \exp\left\{ - \inf_{(x,y) \in \mathcal{S}_2} (x - 2y) \right\} \right]$$

$$\leq \exp\left\{ -1.5s^2 - s^2 (1 - \varepsilon) + 2s^2 (1 - \varepsilon)^2 + \sqrt{n} d_i s + (D_{16}s^3 / \sqrt{n}) \right\}$$

$$\leq \exp\left\{ -\frac{1}{2}s^2 - 2\varepsilon s^2 + \sqrt{n} d_i s + (D_{16}s^3 / \sqrt{n}) \right\}$$

$$\leq \exp\left\{ -\frac{1}{2}(1 + \alpha)s^2 + \sqrt{n} d_i s + (D_{16}s^3 / \sqrt{n}) \right\}.$$

Similarly, we may prove (3.9) for k = 5. This completes the derivation of (3.9) and, hence, also the proof of the lemma.

3.3. Step 3: Blocks and expected numbers of level exceedences

Partition the set of positive integers into small blocks, each of length $\kappa + 1$, where κ is as in (2.2), and large blocks, each of length ℓ , where ℓ is a divergent function of p. We shall take

$$\ell \sim \exp(\frac{1}{4}s^2),\tag{3.11}$$

where $s \to \infty$ as p increases. The integers in each block are consecutive, each consecutive pair of large blocks is separated by a small block and the block furthest to the left is a large block. Let the small blocks be b_1, b_2, \ldots and the large blocks be B_1, B_2, \ldots , indexed such that the order of the blocks is $B_1, b_1, B_2, b_2, \ldots$ Let $B = B_1 = \{1, \ldots, \ell\}$ denote the first large block and let N_1 be the number of indices $i \in B$ for which $R_i > s$. We wish to develop a bound for $E\{N_1I(N_1 \ge 2)\}$. Identical bounds can be derived, uniformly in the block indices, for the versions of N_1 in the case of blocks B_2, B_3, \ldots ; for notational simplicity, we focus solely on B_1 .

By Hölder's inequality,

$$E\{N_1I(N_1 \ge 2)\} \le (EN_1^{a_1})^{1/a_1}P(N_1 \ge 2)^{1/a_2},\tag{3.12}$$

where $a_1, a_2 > 1$ satisfy $a_1^{-1} + a_2^{-1} = 1$. Define $d^0 = \sqrt{n} \max_{1 \le i \le p} d_i$. In view of (3.2) and (3.3),

$$P(N_1 \ge 2) = P(\text{for some } i_1, i_2 \in B \text{ with } i_1 < i_2, R_{i_1}, R_{i_2} > s)$$

$$\le \sum_{i_1=1}^{\ell-1} \sum_{i_2=i_1+1}^{\ell} P(R_{i_1} > s, R_{i_2} > s)$$

$$= \sum_{i_1=1}^{\ell-1} \sum_{i_2=i_1+1}^{\min(i_1+\kappa+1,\ell)} P(R_{i_1} > s, R_{i_2} > s)$$

$$+ \sum_{i_1=1}^{\ell-1} \sum_{i_2=\min(i_1+\kappa+2,\ell)}^{\ell} P(R_{i_1} > s) P(R_{i_2} > s)$$

$$\leq D_{17} \exp(D_{18}s^3 n^{-1/2} + D_{19}d^0 s)$$

$$\times \left[\ell \exp\left\{ -\frac{1}{2}(\alpha+1)s^2 \right\} + \ell^2 \exp(-s^2) \right].$$
(3.13)

Noting that N_1 can be written as $\kappa + 1$ sums of $\ell/(\kappa + 1)$ independent and identically distributed random variables and using calculations based on the binomial distribution, it can be shown that, for the choice of ℓ at (3.11), $E(N_1^{a_1})$ is bounded as $p \to \infty$ for each $a_1 > 0$. Hence, using (3.12) and (3.13), we deduce that for each $\eta_2 \in (0, 1)$,

$$E\{N_1I(N_1 \ge 2)\} \le D_{20} \exp(D_{21}s^3n^{-1/2} + D_{22}d^0s) \times \left[\ell \exp\{-\frac{1}{2}(\alpha+1)s^2\} + \ell^2 \exp(-s^2)\right]^{1-\eta_2}.$$
(3.14)

Write N_2 for the number of exceedences of s that occur in the union of the small blocks b_j that intersect the interval [1, p]. There are $O(p/\ell)$ such small blocks and each is of length $\kappa + 1$, so, by (3.2),

$$E(N_2) \le D_{23}p\ell^{-1}P(R_1 > s) \le D_{24}p\ell^{-1}\exp(D_3s^3n^{-1/2} - \frac{1}{2}s^2 + d^0s).$$
 (3.15)

Provided we choose s = s(p) to diverge to infinity in such a manner that

$$s = O(\sqrt{\log p}), \qquad d^0 = o(s),$$
 (3.16)

it follows from (2.3) that $s^3n^{-1/2} + d^0s = o(s^2)$ and so (3.14) entails that

$$E\{N_1I(N_1 \ge 2)\} = \exp\{\mathrm{o}(s^2)\}[\ell \exp\{-\frac{1}{2}(\alpha+1)s^2\} + \ell^2 \exp(-s^2)]^{1-\eta_2}.$$

Since this is true for each $\eta_2 > 0$, we have

$$E\{N_1I(N_1 \ge 2)\} = \exp\{o(s^2)\} [\ell \exp\{-\frac{1}{2}(\alpha+1)s^2\} + \ell^2 \exp(-s^2)].$$
 (3.17)

3.4. Step 4: Bound for $P(N_1 \ge 1)$, and related bounds

Let N_3 denote the number of exceedences of s which come from large blocks B_j , $1 \le j \le m$, that have two or more exceedences. Write $\sum_j \pi_j$ for the sum over $1 \le j \le m$ of the probability π_j that $R_i > s$ for some $i \in B_j$. Then (a) the expected number of exceedences of s by R_1, \ldots, R_p equals $\sum_{i \le p} P(R_i > s)$ and is less than or equal to $\sum_j \pi_j + E(N_2) + E(N_3)$; (b) the expected number of exceedences in (a) is greater than or equal

to $\sum_{j \leq m-1} \pi_j$; and (c) since $P(N_1 \geq 1) \leq E(N_1) = \ell P(R_1 > s)$ and $P(R_1 > s)$ satisfies (3.2), we have

$$\pi_1 = P(N_1 \ge 1) \le P^0 \equiv D_2 s^{-1} \ell \exp(D_3 s^3 n^{-1/2} - \frac{1}{2} s^2 + d^0 s)$$
 (3.18)

and an identical bound holds for π_1, \ldots, π_m , in particular, (d) $\pi_m \leq P^0$. Results (a)–(d) imply that

$$\left| \sum_{j=1}^{m} \pi_j - \sum_{i=1}^{p} P(R_i > s) \right| \le E(N_2) + E(N_3) + P^0.$$
 (3.19)

Since $E(N_3) \le mE\{N_1I(N_1 \ge 2)\}$, $m = O(p/\ell)$ and bounds for $E\{N_1I(N_1 \ge 2)\}$, $E(N_2)$ and $P(N_1 \ge 1)$ are given by (3.17), (3.15) and (3.18), it follows that (3.19) entails, on taking ℓ as in (3.11),

$$\left| \sum_{j=1}^{m} \pi_j - \sum_{i=1}^{p} P(R_i > s) \right| = \exp\left\{ -\frac{1}{4}s^2 + o(s^2) \right\} \left\{ 1 + p \exp\left(-\frac{1}{4}s^2 - \frac{1}{2}\alpha s^2 \right) \right\}. \quad (3.20)$$

3.5. Step 5: Probabilities of level exceedences

Let \mathcal{F} denote the event that (a) there are no exceedences of s in any of the small blocks that are wholly contained within [1,p]; (b) in each of the large blocks that is wholly contained within [1,p], there is at most one exceedence of s; and (c) there are no exceedences of s in any block fragment that overlaps the end point p. Write \mathcal{G} for the complement of \mathcal{F} . Results (3.15), (3.17) and (3.18) imply that, with ℓ given by (3.11) and assuming that (3.16) holds,

$$P(\mathcal{G}) \le \exp\{-\frac{1}{4}s^2 + o(s^2)\}\{1 + p\exp(-\frac{1}{4}s^2 - \frac{1}{2}\alpha s^2)\}.$$
 (3.21)

Therefore, in order for $P(\mathcal{G}) \to 0$, it is sufficient that for some $\eta_3 \in (0,1)$ and all sufficiently large p, we have

$$(1+\eta_3)\sqrt{2\gamma^{-1}\log p} \le s = \mathcal{O}(\sqrt{\log p}),\tag{3.22}$$

where γ is as defined in Section 2. This choice of s satisfies (3.16) and so if s is given by (3.22), then $P(\mathcal{G})$ satisfies (3.21).

3.6. Step 6: Strong approximation

Let M_j , $1 \le j \le m$, be the number of times that $R_i > s$ for $i \in B_j$. Then, the number, N, say, of blocks B_j for which $M_j \ge 1$ is distributed as $\sum_j I_j$, where the random variables I_j are independent, $I_j = 1$ if $M_j \ge 1$ and $I_j = 0$ otherwise. As before, we define $\pi_j = P(M_j \ge 1)$

1). Conditional on N and on the events " $M_{j_1} \ge 1$ " and " $M_{j_2} \ge 1$," where $1 \le j_1 < j_2 \le m$, the sequences $\{R_i: i \in B_{j_1}\}$ and $\{R_i: i \in B_{j_2}\}$ are independent.

Order the blocks B_j for which $M_j \geq 1$, giving B_{J_1}, \ldots, B_{J_N} , where $1 \leq J_1 < \cdots < J_N \leq m$, and let W_k denote a value of R_i for which $R_i > s$, randomly chosen among such values for which $i \in B_{J_k}$. Write $i = I_k$ for the index of the value of R_i that is chosen as W_k . Then, conditional on N, the random variables W_1, \ldots, W_N are independent and identically distributed as $R(s), J_1, \ldots, J_N$ is a set of integers chosen independently and randomly from $1, \ldots, m$ and I_k is uniformly distributed among indices in B_{J_k} .

Let R'_1, \ldots, R'_p be independent random variables having the distributions of R_1, \ldots, R_p , respectively, let M'_j denote the number of times that R'_i exceeds s for $i \in B_j$ and put $\pi'_j = P(M'_j \ge 1)$. The numbers N' of blocks B_j for which $M'_j \ge 1$ are distributed as $\sum_j I'_j$, where the random variables I'_j are independent and $I'_j = 1$ if $M'_j \ge 1$, $I'_j = 0$ otherwise. An argument similar to, but simpler than, that leading to (3.20) shows that

$$\sum_{j=1}^{m} |\pi_j - \pi_j'| \le \exp\left\{-\frac{1}{4}s^2 + o(s^2)\right\} \left\{1 + p \exp\left(-\frac{1}{4}s^2 - \frac{1}{2}\alpha s^2\right)\right\}. \tag{3.23}$$

By enlarging the probability space if necessary, we can think of N as denoting the number out of m independent and random variables U_1, \ldots, U_j , each uniformly distributed on [0,1], which lie in the respective intervals $[0,\pi_j]$. Take N' to be the number of U_i 's that lie in $[0,\pi'_j]$. Then,

$$P(N = N') \ge 1 - \sum_{j=1}^{m} |\pi_j - \pi'_j|. \tag{3.24}$$

We have already constructed sequences W_1, \ldots, W_N , I_1, \ldots, I_N and J_1, \ldots, J_N . If N' > N, then, conditional on these quantities and on N and N', we select new values $W_{N+1}, \ldots, W_{N'}$, $I_{N+1}, \ldots, I_{N'}$ and $J_{N+1}, \ldots, J_{N'}$ which are independent of W_1, \ldots, W_N , I_1, \ldots, I_N and J_1, \ldots, J_N , with $W_{N+1}, \ldots, W_{N'}$ independently distributed as R(s), the values of $J_{N+1}, \ldots, J_{N'}$ independently and uniformly distributed among $\{1, \ldots, m\} \setminus \{J_1, \ldots, J_N\}$ and the values of $I_{N+1}, \ldots, I_{N'}$ uniformly distributed within the blocks $B_{J_{N+1}}, \ldots, B_{J_{N'}}$, respectively. In this instance, we take $W'_1, \ldots, W'_{N'}$ and $I'_1, \ldots, I'_{N'}$ to be identical to $W_1, \ldots, W_{N'}$ and $I_1, \ldots, I_{N'}$, respectively. If N' < N, then we take $(W'_1, J'_1), \ldots, (W'_{N'}, J'_{N'})$ to be the (exceedence, block index) pairs that remain after randomly and independently deleting N - N' pairs from the sequence $(W_1, J_1), \ldots, (W_N, J_N)$.

Let N_0 denote the number of exceedences of s by R'_1, \ldots, R'_p and let N' represent the number of large blocks B_j in which there is at least one exceedence of s by the sequence R'_1, \ldots, R'_p . Then, $P(N_0 \ge N') = 1$. Conditional on N_0 and N', let $W'_{N'+1}, \ldots, W'_{N_0}$ denote independent and identically random variables, all distributed as R(s), and distribute the locations $I'_{N'+1}, \ldots, I'_{N_0}$ of these exceedences independently and uniformly over the points $\{1, \ldots, p\} \setminus \{I'_1, \ldots, I'_{N'}\}$, conditional on all of the variables N', N_0 , $W'_1, \ldots, W'_{N'}$ and $J'_1, \ldots, J'_{N'}$. Take the values of R'_1, \ldots, R'_p that exceed s to be the variables W'_1, \ldots, W'_{N_0} and let the locations of those exceedences be the points I'_1, \ldots, I'_{N_0} . By

construction, W'_1, \ldots, W'_{N_0} are distributed as the exceedences of s by p independent and identically distributed random variables distributed as R(s); conjointly, I'_1, \ldots, I'_{N_0} are distributed as the locations of those exceedences and the probability that $N_0 = N' = N$, $M_j \in \{0,1\}$ for each $j \in [1,m]$ and there are no exceedences of s in any of the small blocks b_j for any $j \in [1,m]$ is bounded below by $1 - \tau(s)$, where $\tau(s)$ satisfies (2.6); see also (3.20), (3.21), (3.23) and (3.24).

Hence, provided that s satisfies (3.22), we may construct a sequence R'_1, \ldots, R'_p of independent variables with the same marginal distribution as R_1 and such that, with probability bounded below by $1-\tau(s)$, the exceedences of R_1, \ldots, R_p over s are identical to those of R'_1, \ldots, R'_p . The theorem follows from this property, (2.3) and (3.1), on taking s = t.

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